



PTO/SB/08A (08-03)

Substitute for form 1449A INFORMATION DISCLOSURE STATEMENT BY APPLICANT (use as many sheets as necessary)			Complete if Known	
			Application Number	10/678,765
Sheet 1 of 7			Filing Date	October 2, 2003
			First Named Inventor	Serbedzija, George N.
			Art Unit	1632
			Examiner Name	Ram Shulka
			Attorney Docket Number	018852-000511US

U.S. PATENT DOCUMENTS+					
Examiner Initials*	Cite No. ¹	Document Number	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Number Kind Code ² (if known)			
VB	AA	US-6,656,449	12-02-2003	Serbedzija et al.	
	AB	US-6,299,858	10-09-2001	Serbedzija et al.	
	AC	US-5,932,418	08-03-1999	Yager	
	AD	US-5,744,499	04-28-1998	Quash et al.	
	AE	US-5,712,395	01-27-1998	App et al.	
	AF	US-5,681,703	10-28-1997	Tomei	
	AG	US-5,672,470	09-30-1997	Hengstenberg et al.	
	AH	US-5,658,751	08-19-1997	Yue et al.	
	AI	US-5,573,909	11-12-1996	Singer et al.	
	AJ	US-5,565,187	10-15-1996	Zikria et al.	
	AK	US-5,510,099	04-23-1996	Short et al.	
	AL	US-5,506,208	04-09-1996	Eyal et al.	
	AM	US-5,491,084	02-13-1996	Chalfie et al.	
	AN	US-5,424,209	06-13-1995	Kearney	
	AO	US-5,326,692	07-05-1994	Brinkley et al.	
	AP	US-4,816,392	03-28-1989	Hokama	
	AQ	US-4,329,424	05-11-1982	Machlowitz et al.	
	AR	US-4,286,983	09-01-1981	Van Gilse et al.	
VB	AS	US-4,220,858	09-02-1980	Ikeguchi et al.	

FOREIGN PATENT DOCUMENTS								
Examiner Initials*	Cite No. ¹	Foreign Patent Document			Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	T ⁶
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	AX	WO	95/11989	A1	05-04-1995			<input type="checkbox"/>
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VB	AZ	DE	4113393	A1	10-29-1992			abst. only
								<input type="checkbox"/>

Examiner Signature	/Valarie Bertoglio/	Date Considered	07/18/2006
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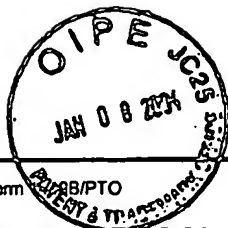
Substitute for form 1449B/PTO			Complete if Known		
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			Art Unit	1632	
			Examiner Name	Ram Shulka	
Sheet	2	of	7	Attorney Docket Number	018852-000511US

NON PATENT LITERATURE DOCUMENTS				
Examiner Initials *	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T ²	
VB	BA	Amsterdam et al., "Requirements for green fluorescent protein detection in transgenic zebrafish embryos", <u>Gene</u> , 173: 99-103 (1996).		
	BB	Armant et al., "Exposure of Embryonic Cells to Alcohol: Contrasting Effects During Preimplantation and Postimplantation Development", <u>Seminars in Perinatology</u> , 20(2): 127-139 (April 1996).		
	BC	Bakkers et al., "An important developmental role for oligosaccharides during early embryogenesis of cyprinid fish", <u>Pro. Natl. Acad. Sci. USA</u> , 94: 7982-7986 (July 1997).		
	BD	Baumann et al., "Bipartite Axiation Follows Incomplete Epiboly in Zebrafish Embryos Treated With Chemical Teratogens", <u>The Journal of Experimental Zoology</u> , 230: 363-376 (1984).		
	BE	Bechter et al., "Teratogenicity of arotinoids (retinoids): comparison of the whole embryo culture system with the in vivo mouse model and the limb bud cell culture assay", <u>Teratol.</u> 44: 29A (1991).		
	BF	Becker et al., "Teratogenic Actions of Ethanol in the Mouse: A Minireview", <u>Pharmacol. Biochem. Behav.</u> , 55(4): 501-513 (1996).		
	BG	Birge et al., "Acetaminophen Hepatotoxicity: Correspondence of Selective Protein Arylation in Human and Mouse Liver <i>in Vitro</i> , in Culture, and <i>in Vivo</i> ", <u>Toxicol. Applied Pharmacol.</u> , 105: 472-482 (1990).		
	BH	Boehm, et al., "Antiangiogenic therapy of experimental cancer does not induce acquired drug resistance", <u>Nature</u> , 390: 404-407 (1997).		
	BI	Bonn, "Blocking angiogenesis in diabetic retinopathy", <u>Lancet</u> , 348: 604 (1996).		
	BJ	Breier et al., "Angiogenesis in Embryos and Ischemic Diseases", <u>Thromb. Haemost.</u> , 78(1): 678-683 (1997).		
	BK	Bresch., "Early Life-Stage Test in Zebrafish versus a Growth Test in Rainbow Trout to Evaluate Toxic Effects", <u>Bull. Environ. Contam. Toxicol.</u> , 46: 641-648 (1991).		
	BL	Buchmann et al., "Immunohistochemical Localization of the Cytochrome P450 Isozymes LMC2 and LM4B (P4501A1) in 2,3,7,8-Tetrachlorodibenzo- <i>p</i> -dioxin Treated Zebrafish (<i>Brachydanio rerio</i>)", <u>Toxicol. Appl. Pharmacol.</u> , 123: 160-169 (1993).		
	BM	Chen et al., "Mutations affecting the cardiovascular system and other internal organs in zebrafish", <u>Development</u> , 123: 293-302 (1996).		
	BN	Chatot et al., "Successful Culture of Rat Embryos on Human Serum: Use in the Detection of Teratogens", <u>Science</u> , 207: 1471-1473 (1980).		
	BO	Chow et al., "The nuclear receptor transcription factor, retinoid-related orphan receptor β , regulates retinal progenitor proliferation", <u>Mech. Dev.</u> , 77(2): 149-164 (1998).		
VB	BP	Cicurel et al., "Post-implantation embryo culture: validation with selected compounds for teratogenicity testing", <u>Xenobiotica</u> , 18(6): 617-624 (1988).		

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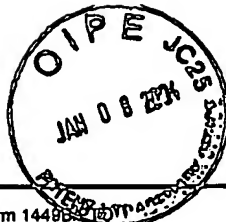
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			Examiner Name	Ram Shulka	
Sheet	3	of	7	Attorney Docket Number	018852-000511US

NON PATENT LITERATURE DOCUMENTS			
Examiner Initials *	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T ²
VB	BQ	Clarke, "Developmental cell death: morphological diversity and multiple mechanisms", <u>Anat. Embryol.</u> , 181: 195-213 (1990).	
	BR	Collodi et al., "Induction of zebrafish (<i>Brachydanio rerio</i>) P450 <i>in vivo</i> and in cell culture", <u>Xenobiotica</u> , 24(6): 487-493 (1994).	
	BS	Couffignal et al., "Animal Model, Mouse Model of Angiogenesis", <u>Am. J. Pathol.</u> , 152(6): 1667-1679 (June 1998).	
	BT	Dawson, "Additive Incidence of Developmental Malformation for <i>Xenopus</i> Embryos Exposed to a Mixture of Ten Aliphatic Carboxylic Acids", <u>Teratology</u> , 44: 531-546 (1991).	
	BU	Debus et al., "Nematode Test to Estimate the Hazard Potential of Solved Contaminations", <u>Chemosphere</u> , 29(3): 611-621 (1994).	
	BV	Drake et al., "Exogenous vascular endothelial growth factor induces malformed and hyperfused vessels during embryonic neovascularization", <u>Proc. Natl. Acad. Sci. U.S.A.</u> , 92: 7657-7661 (August 1995).	
	BW	Driever et al., "Zebrafish: genetic tools for studying vertebrate development," <u>Trends in Genetics</u> , 10:152-159 (1994)	
	BX	Dubois et al., "DNA adducts and P450 induction in human, rat and avian liver cells after exposure to polychlorobiphenyls", <u>Mutation Res.</u> , 345: 181-190 (1995).	
	BY	Dumont et al., "Vascularization of the Mouse Embryo: A Study of <i>flk-1</i> , <i>tek</i> , <i>tie</i> , and Vascular Endothelial Growth Factor Expression During Development", <u>Development Dynamics</u> , 203: 80-92 (1995).	
	BZ	Eisses, "Teratogenicity and Toxicity of Ethylene Glycol Monomethyl Ether (2-Methoxyethanol) in <i>Drosophila melanogaster</i> : Involvement of Alcohol Dehydrogenase Activity", <u>Teratog. Carcinog. Mutagen.</u> , 9: 315-325 (1989).	
	CA	Ellies et al., "Specific craniofacial cartilage dysmorphogenesis coincides with a loss of <i>dlx</i> gene expression in retinoic acid-treated zebrafish embryos", <u>Mechanisms of Development</u> , 61: 23-36 (1997).	
	CB	Ensenbach et al., "Toxicity of Complex Chemical Mixtures: Acute and Long-Term Effects on Different Life Stages of Zebrafish (<i>Brachydanio rerio</i>)", <u>Ecotoxicology and Environmental Safety</u> , 30: 151-157 (1995).	
	CC	Feucht et al., "VEGF Induces Cardiovascular Malformation and Embryonic Lethality", <u>Am. J. Pathology</u> , 151(5):1407-1416 (November 1997).	
	CD	Folkman, "Clinical Applications of Research on Angiogenesis", <u>N. Eng. J. Med.</u> , 333(26): 1757-1763 (December 28, 1995).	
	CE	Fort et al., "Phase III Interlaboratory Study of Fetax, Part 2: Interlaboratory Validation of an Exogenous Metabolic Activation System for Frog Embryo Teratogenesis Assay-Xenopus (Fetax)", <u>Drug and Chem. Toxicol.</u> , 21(1): 1-14 (1998).	
VB	CF	Fouquet et al., "Vessel Patterning in the Embryo of the Zebrafish: Guidance by Notochord", <u>Developmental Biology</u> , 183: 37-48 (1997).	

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Sheet	4	of	7	Attorney Docket Number	018852-000511US

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Examiner Initials *	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T ²	
VB	CG	Frank et al., "Residues of Insecticides, Fungicides, and Herbicides on Ontario-Grown Vegetables, 1980-1985", <u>J. Assoc. Off. Anal. Chem.</u> , 70(6): 1081-1086 (1987).		
	CH	Gavrieli et al., "Identification of Programmed Cell Death In Situ via Specific Labeling of Nuclear DNA Fragmentation", <u>J. Cell Biol.</u> , 119(3): 493-501 (November 1992).		
	CI	Granato et al., "Fishing for genes controlling development", <u>Cur. Op. Gen. Dev.</u> , 6: 461-468 (1996).		
	CJ	Guest et al., "Evaluation of the rat embryo culture system as a predictive test for human teratogens", <u>Can. J. Physiol. Pharmacol.</u> , 72: 57-62 (1994).		
	CK	Hammerschmidt et al., "Genetic analysis of dorsoventral pattern formation in the zebrafish: requirement of a BMP-like ventralizing activity and its dorsal repressor", <u>Genes and Devel.</u> , 10: 2452-2461 (1996).		
	CL	Hanahan, "Signaling Vascular Morphogenesis and Maintenance", <u>Science</u> , 277: 48-50 (July 4, 1997).		
	CM	Heller et al., "Discovery and analysis of inflammatory disease-related genes using cDNA microarrays", <u>Proc. Natl. Acad. Sci. USA</u> , 94: 2150-2155 (March 1997).		
	CN	Herbst et al., "Differential Effects of Laminin, Intact Type IV Collagen, and Specific Domains of Type IV Collagen on Endothelial Cell Adhesion and Migration", <u>J. Cell Biol.</u> , 106: 1365-1373 (April 1988).		
	CO	Hetts, "To Die or Not to Die: An Overview of Apoptosis and its Role in Disease", <u>JAMA</u> , 279(4): 300-307 (January 28, 1998).		
	CP	Hitchcock et al., "Investigations into Using the Nematode <i>Caenorhabditis elegans</i> for Municipal and Industrial Wastewater Toxicity Testing", <u>Arch. Environ. Contam. Toxicol.</u> , 33: 252-260 (1997).		
	CQ	Hunt et al., "A distinct Hox code for the branchial region of the vertebrate head", <u>Nature</u> , 353: 861-864 (October 31, 1991).		
	CR	Iida et al., "Suppression of arachidonic acid cascade-mediated apoptosis in aflatoxin B1-induced rat hepatoma cells by glucocorticoids", <u>Carcinogenesis</u> , 19(7): 1191-1202 (1998).		
	CS	Jain et al., "Quantitative angiogenesis assays: Progress and problems", <u>Nat. Med.</u> , 3(11): 1203-1208 (November 1997).		
	CT	Kerbel, "A cancer therapy resistant to resistance", <u>Nature</u> , 390: 335-336 (November 27, 1997).		
	CU	Kerr et al., "Apoptosis: A Basic Biological Phenomenon With Wide-Ranging Implications in Tissue Kinetics", <u>Br. J. Cancer</u> , 26: 239-257 (1972).		
VB	CV	Kim et al., "Requirement for Specific Proteases in Cancer Cell Intravasation as Revealed by a Novel Semiquantitative PCR-Based Assay", <u>Cell</u> , 94:353-362 (1998).		

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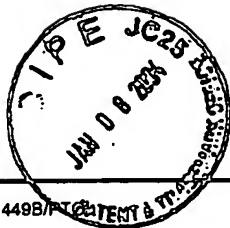
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VB	CW	Knight, "Adverse Drug Reactions in Neonates", <u>J. Clin. Pharmacol.</u> 34: 128-135 (1994).	
	CX	Law et al., "Hepatotoxicity of the drinking water disinfection by-product, dichloroacetic acid, in the medaka small fish model", <u>Toxicol. Letters</u> , 94: 19-27 (1998).	
	CY	Liao et al., "The zebrafish gene <i>cloche</i> acts upstream of a <i>flk-1</i> homologue to regulate endothelial cell differentiation", <u>Development</u> , 124: 381-389 (1997).	
	CZ	Liepins et al., "Cell Injury and Apoptosis", <u>Scanning Microscopy</u> , 8(3): 631-643 (1994).	
	DA	Long et al., "GATA-1 expression pattern can be recapitulated in living transgenic zebrafish using GFP reporter gene", <u>Development</u> , 124: 4105-4111 (1997).	
	DB	Losordo et al., "Gene Therapy for Myocardial Angiogenesis: Initial Clinical Results With Direct Myocardial Injection of phVEGF ₁₆₅ as Sole Therapy for Myocardial Ischemia", <u>Circulation</u> , 98: 2800-2804 (1998).	
	DC	Marret et al., "Caffeine-induced disturbances of early neurogenesis in whole mouse embryo cultures", <u>Brain Research</u> , 773: 213-216 (1997).	
	DD	McDanell et al., "Effect of Dietary Fat on the <i>In Vitro</i> Hepatotoxicity of Paracetamol", <u>Biochemical Pharmacology</u> , 44(7): 1303-1306 (1992).	
	DE	Mizell et al., "The aquatic vertebrate embryo as a sentinel for toxins: zebrafish embryo dechorionation and perivitelline space microinjection", <u>Int. J. Dev. Biol.</u> , 41: 411-423 (1997).	
	DF	Monteith et al., "Comparison of Tacrine-Induced Cytotoxicity in Primary Cultures of Rat, Mouse, Monkey, Dog, Rabbit, and Human Hepatocytes", <u>Drug and Chem. Toxicol.</u> , 19(1&2): 59-70 (1996).	
	DG	Muller et al., "Effect of Concentration on the Cytotoxic Mechanism of Doxorubicin-Apoptosis and Oxidative DNA Damage", <u>Biochem. Biophys. Res. Comm.</u> , 230(2): 254-257 (1997).	
	DH	Mundle et al., "Two <i>in Situ</i> Labeling Techniques Reveal Different Patterns of DNA Fragmentation during Spontaneous Apoptosis <i>In Vivo</i> and Induced Apoptosis <i>In Vitro</i> ", <u>Anticancer Res.</u> , 15: 1895-1904 (1995).	
	DI	O'Reilly et al., "Endostatin: An Endogenous Inhibitor of Angiogenesis and Tumor Growth", <u>Cell</u> , 88: 277-285 (January 24, 1997).	
	DJ	Ozato et al., "Production of transgenic fish: introduction and expression of chicken δ -crystallin gene in medaka embryos", <u>Cell Differentiation</u> , 19:237-244 (1986).	
	DK	Relou et al., "Effect of culture conditions on endothelial cell growth and responsiveness", <u>Tissue & Cell</u> , 30(5): 525-530 (1998).	
VB	DL	Rivard et al., "Age-Dependent Impairment of Angiogenesis", <u>Circulation</u> , 99: 111-120 (1999).	

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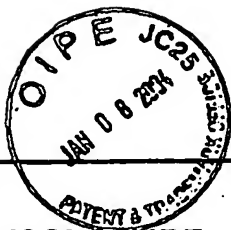
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VB	DM	Roebuck et al., "A Review of the Neuroanatomical Findings in Children with Fetal Alcohol Syndrome or Prenatal Exposure to Alcohol", <u>Alcoholism: Clin. Exper. Res.</u> , 22(2): 339-344 (April 1998).	
	DN	Serbedzija et al., "Regulation in the heart field of zebrafish", <u>Development</u> , 125: 1095-1101 (1998).	
	DO	Serbedzija et al., "Analysis of Neural Crest Cell Migration in Splotch Mice Using a Neural Crest-Specific LacZ Reporter", <u>Developmental Biology</u> , 185: 139-147 (1997).	
	DP	Serbedzija et al., "Cell Death in the CNS of the Wnt-1 Mutant Mouse", <u>J. Neurobio.</u> , 31(3): 275-282 (1996).	
	DQ	Serbedzija et al., "Vital dye analysis of cranial neural crest cell migration in the mouse embryo", <u>Development</u> , 116: 297-307 (1992).	
	DR	Serbedzija et al., "Developmental potential of trunk neural crest cells in the mouse", <u>Development</u> , 120: 1709-1718 (1994).	
	DS	Semino et al., "Synthesis of 'Nod'-like chitin oligosaccharides by the <i>Xenopus</i> developmental protein DG42", <u>Proc. Natl. Acad. Sci. USA</u> , 92: 3498-3501 (April 1995).	
	DT	Semino et al., "Expression of <i>Rhizobium</i> Chitin Oligosaccharide Fucosyltransferase in Zebrafish Embryos Disrupts Normal Development", <u>Annals New York Academy of Sciences</u> , 842: 49-54 (1998).	
	DU	Semino, C. et al., "Homologs of the <i>Xenopus</i> developmental gene DG-42 are present in zebrafish and mouse and are involved in the synthesis of Nod-like chitin oligosaccharides during early embryogenesis", <u>Proc. Natl. Acad. Sci. USA</u> , 93: 4548-4553 (May 1996).	
	DV	Singer, "Sensitive Fluorescent Stains for Detecting Nucleic Acids in Gels and Solutions", <u>Biotechnol. Intl.</u> , 1: 267-276 (1997).	
	DW	Stainier et al., "Mutations affecting the formation and function of the cardiovascular system in the zebrafish embryo", <u>Development</u> , 123: 285-292 (1996).	
	DX	Stainier et al., "The Zebrafish as a Model System to Study Cardiovascular Development", <u>Trends Cardiovasc. Med.</u> , 4(5): 207-212 (1994).	
	DY	Steller, "Mechanisms and Genes of Cellular Suicide", <u>Science</u> , 267: 1445-1449 (March 10, 1995).	
	DZ	Stringer et al., "Rapid Measurement of Toxicity Using Electrochromic Dyes and Frog Embryos", <u>Bull. Environ. Contam. Toxicol.</u> , 51: 557-563 (1993).	
	EA	Thompson, C.B., "Apoptosis in the Pathogenesis and Treatment of Disease", <u>Science</u> , 267: 1456-1462 (March 10, 1995).	
VB	EB	Thompson, M.A. et al., "The <i>cloche</i> and <i>spadetail</i> Genes Differentially Affect Hematopoiesis and Vasculogenesis", <u>Dev. Biol.</u> , 197: 248-269 (1998).	

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Examiner Initials *	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T ²
VB	EC	Turk et al., "Synthetic Analogues of TNP-470 and Ovalicin Reveal a Common Molecular Basis for Inhibition of Angiogenesis and Immunosuppression", <u>Bioorganic & Medicinal Chemistry</u> , 6: 1163-1169 (1998).	
	ED	Ungvary et al., "Combined Embryotoxic Action of Toluene, a Widely Used Industrial Chemical, and Acetylsalicylic Acid (Aspirin)", <u>Teratology</u> , 27: 261-269 (1983).	
	EE	Van Leeuwen et al., "Fish Embryos as Teratogenicity Screens: A Comparison of Embryotoxicity between Fish and Birds", <u>Ecotoxicol. Environ. Safety</u> , 20: 42-52 (1990).	
	EF	Weinstein et al., " <u>gridlock</u> , a localized heritable vascular patterning defect in the zebrafish", <u>Nature Med.</u> , 1(11): 1143-1147 (November 1995).	
	EG	Wolff et al., "Dexamethasone Increases Hepatotoxicity of MTX in Children with Brain Tumors," <u>Anticancer Res.</u> , 18: 2895-2900 (1998).	
VB	EH	Zetter, "Angiogenesis and Tumor Metastasis", <u>Annual Rev. Med.</u> , 49: 407-424 (1998).	

Examiner Signature	/Valarie Bertoglio/	Date Considered	07/18/2006
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* EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

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